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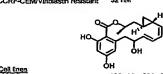
(54) Title: NOVEL MA CROCYCLES AND USES THEREOF

(57) Abstract: The present invention relates to compounds having the structure (and pharmaceutically acceptable derivatives thereof) (I) wherein R,,-R4, Z, X, A-B, D-E, G-J, and K-L are as defined herein, the synthesis thereof and the use of these compounds as therapeutic agents.

IC50 of Growth Inhibition of Different Tumor Cell lines

MCF-7 CCRF-CEM CCRF-CEM/Vinblastin resistant 54 nM 42 nM 32 nM >500 nM

>500 nM



HO H' OH



Cell lines
MCF-7
CCRF-CEM
CCRF-CEM/Vinblastin resistant

150 nM, >500 nM 208 nM 209 nM

390 nM

>500 nM



HOTOLOGIC

Cell lines MCF-7

88 nM

282 nM

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### NOVEL MACROCYCLES AND USES THEREOF

#### ABSTRACT

The present invention relates to compounds having the structure (and pharmaceutically acceptable derivatives thereof)

$$R_3$$
 $R_2$ 
 $R_1$ 
 $R_2$ 
 $R_3$ 
 $R_4$ 
 $R_2$ 
 $R_4$ 
 $R_4$ 
 $R_5$ 
 $R_6$ 
 $R_7$ 
 $R_8$ 
 $R_9$ 
 $R_9$ 

wherein R<sub>0</sub>-R<sub>4</sub>, Z, X, A-B, D-E, G-J, and K-L are as defined herein, the synthesis thereof and the use of these compounds as therapeutic agents.

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